

by estrogen, and inhibits chromatid separation. (Pei, L., and Melmed, S., *Isolation and characterization of a pituitary tumor transforming gene*, Mol. Endocrinol. 11:433-441 [1997]; Zhang, X., *et al.*, *Structure, expression, and function of human pituitary tumor-transforming gene (PTTG)*, Mol. Endocrinol. 13:156-166 [1999a]; Heaney, A.P., *et al.*, *Early involvement of estrogen-induced pituitary tumor transforming gene and fibroblast growth factor expression in prolactinoma pathogenesis*, Nature Med. 5:1317-1321 [1999]; Zou, H., *et al.*, *Identification of a vertebrate sister-chromatid separation inhibitor involved in transformation and tumorigenesis*, Science 285:418-422 [1999]).--.

At page 5, lines 15-25, please delete the entire paragraph, and insert therefor the following paragraph:

--The sequence of events in angiogenesis leading to formation of new blood vessels from pre-existing vessels is highly regulated (Jain, RK *et al.*, *Quantitative angiogenesis assays: progress and problems*, Nat Med. 3:1203-1208 [1997]; Darland DC and D'Amore PA, *Blood vessel maturation: vascular development comes of age*, J Clin Invest. 103:157-158 [1999]), and involves dissolution of vessel basement membranes, and formation of new lumen and pericytes by vascular endothelial cells. During tumor-associated angiogenesis, sustained production of angiogenic factors by cancer cells, or indirect macrophage stimulation, causes disregulated immature vessel growth (Folkman, J. and Shing, Y., *Angiogenesis*, J Biol Chem. 267:10931-10934[1992]). A number of in vitro and in vivo assays have been useful for studying angiogenesis (e.g., Jain, RK *et al.* [1997]; Auerbach, R. *et al.*, *Assays for angiogenesis: a review*, Pharmacol Ther. 51:1-11 [1991]).--.

At page 9, lines 4-7, please delete the entire paragraph, and insert therefor the following paragraph:

--Figure 5 shows *PTTG* mRNA expression in normal adult human T-cells treated with mitogen anti-CD3 antibody. T-cells were isolated and stimulated with anti-CD3 antibody for 72 hours. *PTTG* mRNA was measured with northern blotting (middle panel) and percentage of cells in S or G2/M phase was determined by FACS.--.

At page 9, lines 8-11, please delete the entire paragraph, and insert therefor the following paragraph:

at --Figure 6 shows *PTTG* mRNA expression in normal adult human T-cells treated with mitogen phytohemagglutinin (PHA). T-cells were isolated and stimulated with increasing concentrations of PHA for 72 hours. *PTTG* mRNA was measured with northern blotting (middle panel) and percentage of cells in S or G2/M phase was determined by FACS.--

At page 9, lines 12-15, please delete the entire paragraph, and insert therefor the following paragraph:

at --Figure 7 shows *IL-2* mRNA expression in normal adult human T cells treated with mitogen anti-CD3 antibody. T-cells were isolated and stimulated with anti-CD3 antibody for 72 hours. *IL-2* mRNA (middle panel) was measured with northern blotting and percentage of cells in S or G2/M phase was determined by FACS.--

At page 9, lines 16-18, please delete the entire paragraph, and insert therefor the following paragraph:

at --Figure 8 shows cyclophilin mRNA expression in normal adult human T cells treated with mitogen anti-CD3 antibody. T cells were isolated and stimulated with anti-CD3 antibody for 48 hours. Cyclophilin mRNA (middle panel) was measured with northern blotting.--

At page 9, lines 19-22, please delete the entire paragraph, and insert therefor the following paragraph:

at --Figure 9 demonstrates *PTTG* mRNA expression and hydrocortisone. PHA (5 µg/mL)-stimulated normal adult human T cells were treated with hydrocortisone for 72 hours. *PTTG* mRNA was measured with northern blotting (middle panel) and percentage of cells in S or G2/M phase was determined by FACS.--

At page 9, lines 23-25, please delete the entire paragraph, and insert therefor the following paragraph:

--Figure 10 shows *PTTG* mRNA expression and cyclosporin. PHA (5 $\mu\text{g/mL}$)-stimulated normal adult human T cells were treated with cyclosporin for 72 h. *PTTG* mRNA was measured with northern blotting (middle panel) and percentage of cells in S or G2/M phase was determined by FACS.--

At page 10, lines 4-9, please delete the entire paragraph, and insert therefor the following paragraph:

--Figure 13 shows *PTTG* mRNA expression in human Jurkat T cell leukemia line. Jurkat T cells were treated as described below. (1) cells kept for 48 h in 1% FBS-supplemented culture medium; (2) cells after medium change for fresh 1% FBS-supplemented; (3) cells after medium change for 10% FBS-supplemented; (4) phytohemagglutinin (PHA; 1 $\mu\text{g/mL}$) + phorbol-12-meristate-13-acetate (PMA; 50 ng/mL) in 1% FBS; (5) (PHA + PMA) + cyclosporine A (1 $\mu\text{g/mL}$); (6) (PHA + PMA) + TGF- β 1 (10 ng/mL). *PTTG* mRNA was measured with northern blotting (middle panel) and percentage of cells in S phase was determined by FACS.--

At page 98, lines 13-24, please delete the entire paragraph, and insert therefor the following paragraph:

--Tube forming assay. Matrigel is useful for studying HUVEC attachment and differentiation. Since Matrigel itself induces HUVEC differential activity, we used GFR Matrigel to reduce the effect of growth factors from the Matrigel itself. As shown in Figure 22A, when HUVECs adhered on GFR Matrigel, they aligned with one another and formed tubes resembling a capillary plexus under the influence of differential activity in the CM. Quantitative analysis of HUVEC tube formation (Denekamp J., *Review article: angiogenesis, neovascular proliferation and vascular pathophysiology as targets for cancer therapy*, Br J Radiol. 66:181-196 [1993]) revealed that WT-hPTTG-CM enhanced HUVEC tube formation compared to that observed

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when HUVECs were incubated in CM derived from other cell lines (Figure 22B; $p < 0.01$). The morphologic changes resembling capillary formation were suppressed by adding anti-bFGF antibody to each CM. Suppressive effects of anti-bFGF antibody of WT-hPTTG-CM, M-hPTTG-CM, C-CM and N-CM were, in the same order, 74%, 58%, 57%, and 62%.--

At page 99, lines 1-13, please delete the entire paragraph, and insert therefor the following paragraph:

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